(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization International Bureau





(43) International Publication Date 13 May 2004 (13.05.2004)

PCT

(10) International Publication Number WO 2004/039957 A2

(51) International Patent Classification7:

C12N

(74) Agent: MANDRAGOURAS, Amy, E.; Lahive & Cock-

(21) International Application Number:

PCT/US2003/034424

(22) International Filing Date: 29 October 2003 (29.10.2003)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

60/422,419

29 October 2002 (29.10.2002) US

(71) Applicant (for all designated States except US): THE CENTER FOR BLOOD RESEARCH, INC. [US/US]: 800 Huntington Avenue, Boston, MA 02115 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): LIEBERMAN, Judy [US/US]; 63 Buckminster Road, Brookline, MA (US). SHANKAR, Premlata [-/US]; 46 Hilburn Street. Roslindale, MA 02131 (US), NARASIMHASWAMY, Manjunath [-/US]; 46 Hilburn Street, Roslindale, MA 02131 (US). SONG, Erwei [--/US]; 11 Monument Way, Quincy, MA (US). LEE, Sang-Kyung [—/US]; ** (**).

fiele, LLP, 28 State Street, Boston, MA 02109 (US).

(81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.

(84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

without international search report and to be republished upon receipt of that report

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: INHIBITION OF GENE EXPRESSION USING RNA INTERFERING AGENTS

(57) Abstract: The present invention is based, at least in part, on the discovery of compositions and methods for the treatment and prevention of infectious diseases or disorders, e.g., HIV infection, AIDS, and AIDS-related diseases. In particular, the present invention pertains to methods of modulating cellular gene expression or protein activity, e.g., CCR5, gene expression or protein activity and/or gene expression or protein activity of a gene or sequence of an infectious agent, in order to treat or prevent infectious diseases or disorders, HIV infection, AIDS, or an AIDS-related disease or disorder. In one embodiment, the combination of an RNA interfering agent targeting a cellular gene in combination with an RNA interfering agent targeting a gene or sequence of an infectious agent results in prolonged prevention of infection by an infectious agent. The present invention is based on the identification of novel RNA interference agents, e.g., siRNA molecules, which target cellular genes, e.g., chemokine receptors, e.g., the CCR5 gene, and result in inhibition of target gene expression on target gene expressing cells, thereby inhibiting entry of infectious agents, e.g., HIV infection into target cells, prevention infection, and/or suppressing replication in established infection.

